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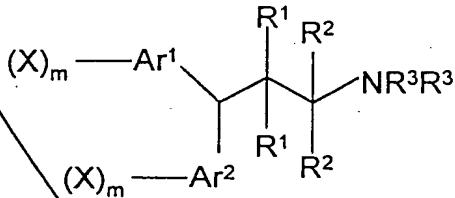
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Claims

*Sub C1* 1. A method of treating a <sup>patient</sup> ~~patient~~ for depression comprising the step of administering to said patient an effective amount of a compound having a NMDA  $IC_{50}$  of about 50 nM to about 1  $\mu$ M as measured in the NMDA assay and a serotonin reuptake  $IC_{50}$  of less than or equal to about 100 nm as measured in the serotonin reuptake inhibition assay. *250 17863*

10 2. The method of claim 1, wherein said compound has an NMDA receptor  $IC_{50}$  of 50 nM to 1  $\mu$ M and a SSRI  $IC_{50}$  less than 100 nM.

*SLB* 15 3. A method of treating a patient for depression comprising the step of administering to said patient an effective amount of a compound having the chemical structure:



wherein each X is independently selected from the group consisting of -Br, -Cl, -F, -I, -CF<sub>3</sub>, alkyl, -OH, -OCF<sub>3</sub>, -O-alkyl, and -O-acyl;

20 Ar<sup>1</sup> and Ar<sup>2</sup> are each independently selected from the group consisting of phenyl, naphthyl, thiofuranyl, tetrahydronaphthyl, furanyl, tetrahydrofuranyl, pyridyl, quinolinyl, isoquinolinyl, tetrahydroquinolinyl, tetrahydroisoquinolinyl, cyclohexyl, cycloheptyl, and cyclopentyl;

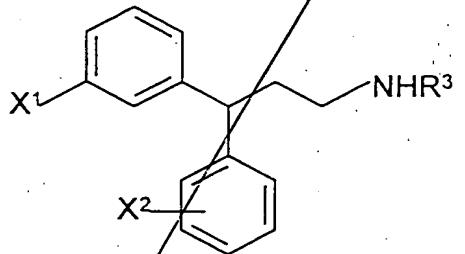
25 each R<sup>1</sup> is independently selected from the group consisting of -H, alkyl, hydroxyalkyl, -OH, -O-alkyl, and -O-acyl;

each R<sup>2</sup> is independently selected from the group consisting of -H, alkyl, and hydroxyalkyl, or both R<sup>2</sup>'s together are imino;

132  
 each  $R^3$  is independently selected from the group consisting of -H, alkyl, 2-hydroxyethyl, and alkylphenyl; and  
 each  $m$  is independently an integer from 0 to 5;  
 provided that if both  $R_3$ 's are -CH<sub>3</sub>, then both  $X_m$ 's are not  
 5 3-F, 4-F, 3-CF<sub>3</sub>, 4-Cl, and if both  $R_3$ 's are -CH<sub>3</sub> and one  $X_m$  is 4-F  
 then the other  $X_m$  is not 4-Cl; further provided that if one  $R_3$  is  
 -H and the other  $R_3$  is -CH<sub>3</sub>, then both  $X_m$ 's are not 4-Cl, and if  
 one  $R_3$  is -H and the other  $R_3$  is -CH<sub>3</sub>, then at least one  $m$  is 1;  
 or a pharmaceutically acceptable salt thereof.

10  
 4. The method of claim 3 wherein for said compound each  $X$   
 is independently either -F, -Cl, -OCF<sub>3</sub>, or -CF<sub>3</sub>;  
 each  $R^1$  is -H;  
 each  $R^2$  is -H;  
 15 one  $R^3$  is -H, and the other  $R^3$  is either -H or -CH<sub>3</sub>; and  
 each  $m$  is 1.

5. The method of claim 3 wherein said compound has the  
 chemical structure:



20  
 wherein  $X^1$  is either -Br, -Cl, -F, -I, -CF<sub>3</sub>, alkyl, -OH,  
 -OCF<sub>3</sub>, -O-alkyl, or -O-acyl.

25  $X^2$  is either -Br, -Cl, -F, -I, -CF<sub>3</sub>, alkyl, -OH, -OCF<sub>3</sub>,  
 -O-alkyl, or -O-acyl; and

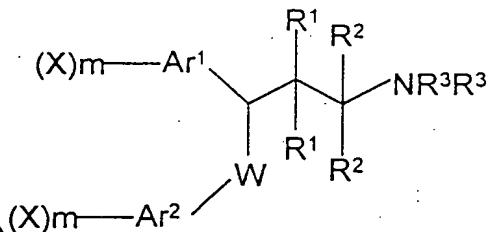
$R^3$  is either -H or -CH<sub>3</sub>;

or a pharmaceutically acceptable salt thereof.

6. The method of claim 5, wherein  $X^1$  is -F, -Cl, -OCF<sub>3</sub>, or  
 30 -CF<sub>3</sub>; and  $X^2$  is either 2-OCH<sub>3</sub>, 2-CH<sub>3</sub>, 3-F, 3-CF<sub>3</sub>, or 4-CF<sub>3</sub>.

*54b*  
*BB*

7. A method of treating a patient for depression comprising the step of administering to said patient an effect amount of a compound having the chemical structure:



5

wherein each  $\text{X}$  is independently selected from the group consisting of  $-\text{Br}$ ,  $-\text{Cl}$ ,  $-\text{F}$ ,  $-\text{I}$ ,  $-\text{CF}_3$ , alkyl,  $-\text{OH}$ ,  $-\text{OCF}_3$ ,  $-\text{O-alkyl}$ , and  $-\text{O-acyl}$ ;

10  $\text{Ar}^1$  and  $\text{Ar}^2$  are each independently selected from the group consisting of phenyl, naphthyl, thiofuranyl, tetrahydronaphthyl, furanyl, tetrahydrofuranyl, pyridyl, quinolinyl, isoquinolinyl, tetrahydroquinolinyl, tetrahydroisoquinolinyl, cyclohexyl, cycloheptyl, and cyclopentyl;

15 each  $\text{R}^1$  is independently selected from the group consisting of  $-\text{H}$ , alkyl, hydroxyalkyl,  $-\text{OH}$ ,  $-\text{O-alkyl}$ , and  $-\text{O-acyl}$ ;

each  $\text{R}^2$  is independently selected from the group consisting of  $-\text{H}$ , alkyl, and hydroxyalkyl, or both  $\text{R}^2$ 's together are imino;

each  $\text{R}^3$  is independently selected from the group consisting of  $-\text{H}$ , alkyl, 2-hydroxyethyl, and alkylphenyl; and

20  $\text{m}$  is 0 to 5;

or a pharmaceutically acceptable salt thereof.

*John C.*  
8. The method of claim 7, wherein for said compound each  $\text{X}$  is independently either  $-\text{F}$ ,  $-\text{Cl}$ ,  $-\text{OCF}_3$  or  $-\text{CF}_3$ ;

25  $\text{Ar}^1$  and  $\text{Ar}^2$  are each independently phenyl or naphthyl;

each  $\text{R}^1$  is  $-\text{H}$ ;

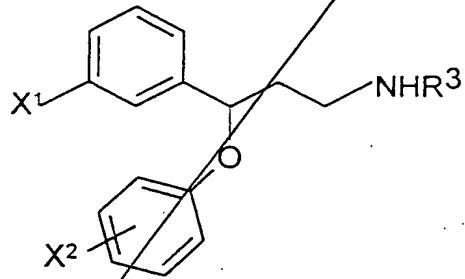
each  $\text{R}^2$  is  $-\text{H}$ ;

one  $\text{R}^3$  is  $-\text{H}$ , and the other  $\text{R}^3$  is either  $-\text{H}$  or  $-\text{CH}_3$ ;

each  $\text{m}$  is 0 or 1.

*Hand C1*

9. The method of claim 7, wherein said compound has the chemical structure:



5 wherein  $\text{X}^1$  is either -Br, -Cl, -F, -I, -CF<sub>3</sub>, alkyl, -OH, -OCF<sub>3</sub>, -O-alkyl, or -O-acyl;

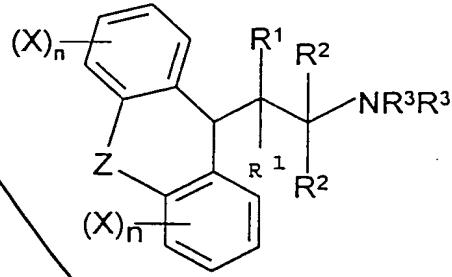
$\text{X}^2$  is either -Br, -Cl, -F, -I, -CF<sub>3</sub>, alkyl, -OH, -OCF<sub>3</sub>, -O-alkyl, or -O-acyl; and

$\text{R}^3$  is either -H or -CH<sub>3</sub>;

10 or a pharmaceutically acceptable salt thereof.

10. The method of claim 9 wherein  $\text{X}^1$  is either -F, -Cl, -OCF<sub>3</sub> or -CF<sub>3</sub>; and  $\text{X}^2$  is either 2-OCH<sub>3</sub>, 2-CH<sub>3</sub>, 3-F, 3-CF<sub>3</sub>, or 4-CF<sub>3</sub>.

*Sub 4*  
15 11. A method of treating a patient for depression comprising the step of administering to said patient an effective amount of a compound having the chemical structure:



20 wherein each X is independently selected from the group consisting of -Br, -Cl, -F, -I, -CF<sub>3</sub>, alkyl, -OH, -OCF<sub>3</sub>, -O-alkyl, and -O-acyl;

34 each  $R^1$  is independently selected from the group consisting of -H, alkyl, hydroxyalkyl, -OH, -O-alkyl, and -O-acyl;

each  $R^2$  is independently selected from the group consisting of -H, alkyl, and hydroxyalkyl, or both  $R^2$ 's together are imino;

5 each  $R^3$  is independently selected from the group consisting of -H, alkyl, 2-hydroxyethyl, and alkylphenyl;

$Z$  is either  $-\text{CH}_2\text{CH}_2-$ ,  $-\text{CH}_2\text{CH}(\text{CH}_3)-$ ,  $-\text{CH}=\text{CH}-$ ,  $-\text{O}-\text{CH}_2-$ ,  $-\text{S}-\text{CH}_2-$ ,  $-\text{CH}_2-$ ,  $-\text{O}-$ , or  $-\text{S}-$ ; and

10 each  $n$  is independently 1 to 4; or a pharmaceutically acceptable salt thereof.

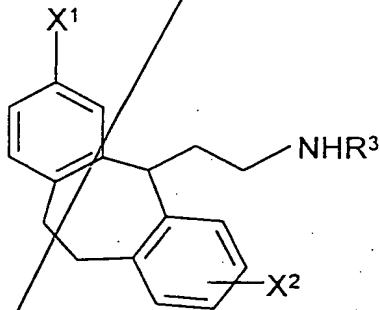
12. The compound of claim 11, wherein each  $X$  is independently either -F, -Cl, -OCF<sub>3</sub>, or -CF<sub>3</sub>;

each  $R^1$  is -H;

each  $R^2$  is -H;

15 one  $R^3$  is -H, and the other  $R^3$  is either -H or -CH; and each  $n$  is 1.

13. The method of claim 11, wherein said compound has the chemical structure:



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wherein  $X^1$  is either -Br, -Cl, -F, -I, -CF<sub>3</sub>, alkyl, -OH, -OCF<sub>3</sub>, -O-alkyl, or -O-acyl;

$X^2$  is either -Br, -Cl, -F, -I, -CF<sub>3</sub>, alkyl, -OH, -OCF<sub>3</sub>, -O-alkyl, or -O-acyl; and

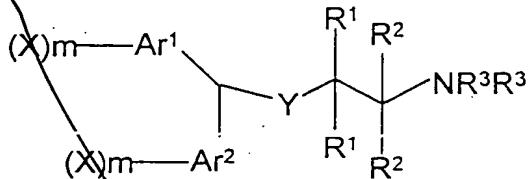
25  $R^3$  is either -H or -CH<sub>3</sub>;

or a pharmaceutically acceptable salt thereof.

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*135*

14. The method of claim 13 wherein  $X^1$  is  $-F$ ,  $-Cl$ ,  $-OCF_3$ , or  $-CF_3$ ; and  $X^2$  is either either  $-F$ ,  $-Cl$ ,  $-OCH_3$ ,  $-CH_3$ ,  $-OCF_3$  or  $-CF_3$ .

5 15. A method of treating a patient for depression comprising the step of administering to said patient an effect amount of a compound having the chemical structure:



10 wherein each  $X$  is independently selected from the group consisting of  $-Br$ ,  $-Cl$ ,  $-F$ ,  $-I$ ,  $-CF_3$ , alkyl,  $-OH$ ,  $-OCF_3$ ,  $-O$ -alkyl, and  $-O$ -acyl; preferably, each  $X$  is independently either  $-F$ ,  $-Cl$ ,  $-OCF_3$  or  $-CF_3$ ;

15  $Ar^1$  and  $Ar^2$  are each independently selected from the group consisting of phenyl, naphthyl, thiofuranyl, tetrahydronaphthyl, furanyl, tetrahydrofuranyl, pyridyl, quinolinyl, isoquinolinyl, tetrahydroquinolinyl, tetrahydroisoquinolinyl, cyclohexyl, cycloheptyl, and cyclopentyl; preferably  $Ar^1$  and  $Ar^2$  are independently naphthyl or phenyl; more preferably at least one of 20  $Ar^1$  and  $Ar^2$  is phenyl; and more preferably, both  $Ar^1$  and  $Ar^2$  are phenyl;

Y is either  $-CH_2-$ ,  $-O-$ , or  $-S-$ ;

25 each  $R^1$  is independently selected from the group consisting of  $-H$ , alkyl, hydroxyalkyl,  $-OH$ ,  $-O$ -alkyl, and  $-O$ -acyl; preferably, each  $R^1$  is  $-H$ ;

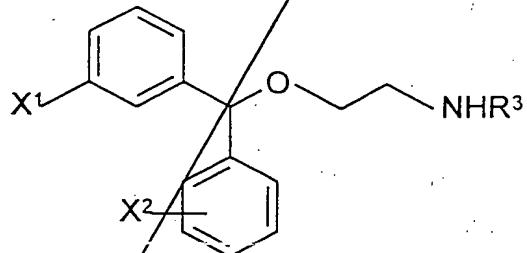
each  $R^2$  is independently selected from the group consisting of  $-H$ , alkyl, and hydroxyalkyl, or both  $R^2$ 's together are imino; preferably each  $R^2$  is  $-H$ ;

30 each  $R^3$  is independently selected from the group consisting of  $-H$ , alkyl, 2-hydroxyethyl, and alkylphenyl; preferably, each  $R^3$  is independently either  $-H$  or  $-CH_3$ ; more preferably one  $R^3$  is  $-H$ , and the other  $R^3$  is either  $-H$  or  $-CH_3$ ; and

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each  $m$  is independently an integer from 0 to 5; and preferably, each  $m$  is independently 0 or 1.

16. The method of claim 15, wherein said compound has the chemical structure; Structure VIII



wherein  $X^1$  is independently selected from the group consisting of -H, -Br, -Cl, -F, -I, -CF<sub>3</sub>, alkyl, -OH, -OCF<sub>3</sub>, -O-alkyl, or -O-acyl; preferably,  $X^1$  is either -F, -Cl, -OCF<sub>3</sub> and -CF<sub>3</sub>;

$X^2$  is either -Br, -Cl, -F, -I, -CF<sub>3</sub>, alkyl, -OH, -OCF<sub>3</sub>, -O-alkyl, or -O-acyl; preferably,  $X^2$  is independently either -F, -Cl, -OCH<sub>3</sub>, -CH<sub>3</sub>, -OCF<sub>3</sub> or -CF<sub>3</sub>; more preferably,  $X^2$  is either 2-OCH<sub>3</sub>, 2-CH<sub>3</sub>, 3-F, 3-CF<sub>3</sub>, or 4-CF<sub>3</sub>; and

$R^3$  is either -H or CH<sub>3</sub>;

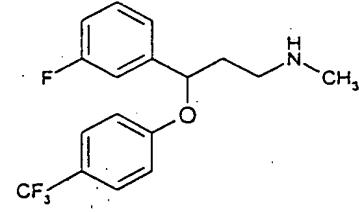
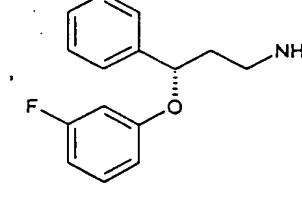
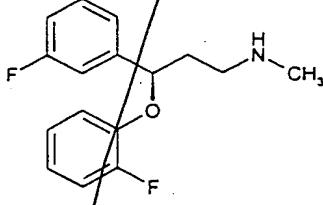
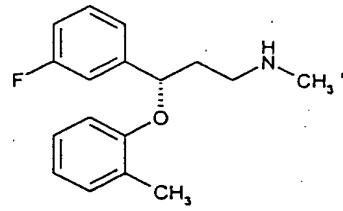
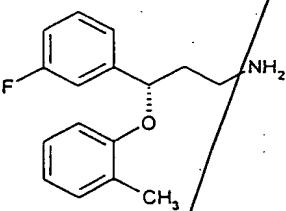
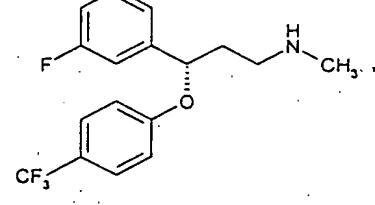
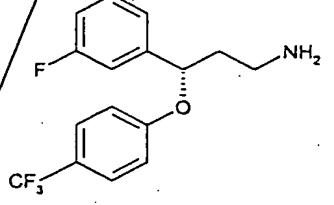
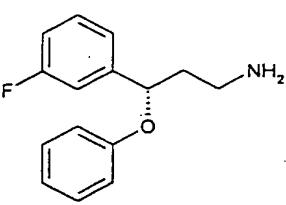
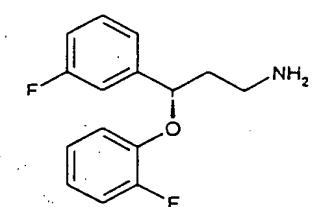
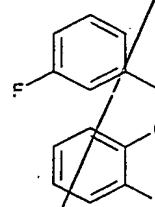
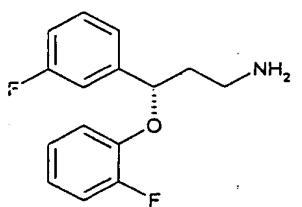
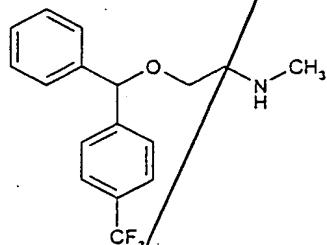
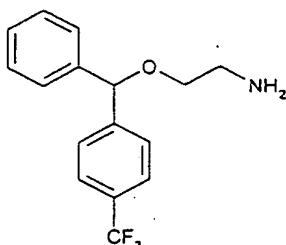
or a pharmaceutically acceptable salt thereof.

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17. A compound having the chemical structure;



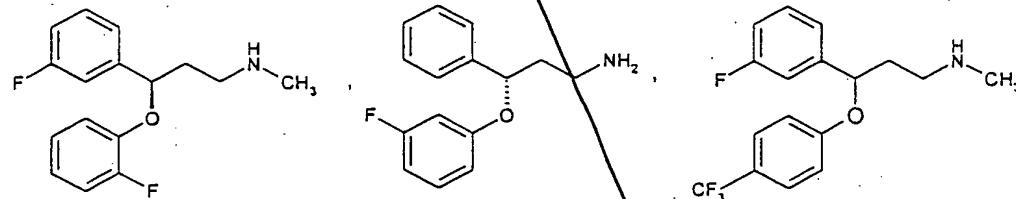
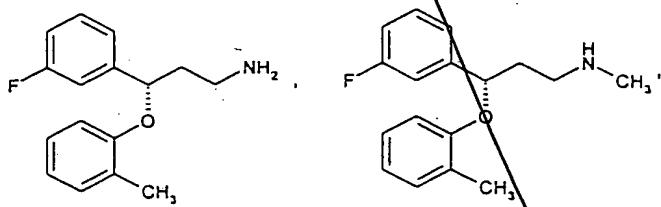
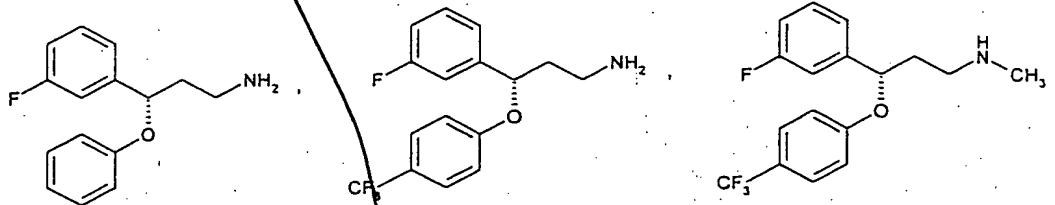
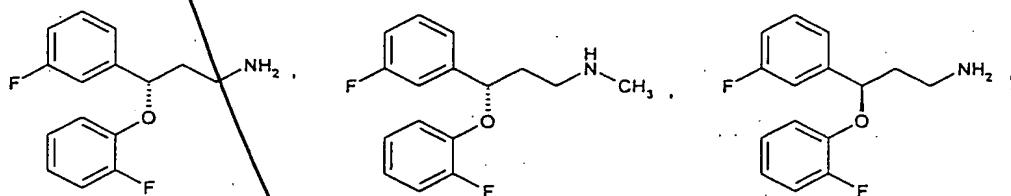
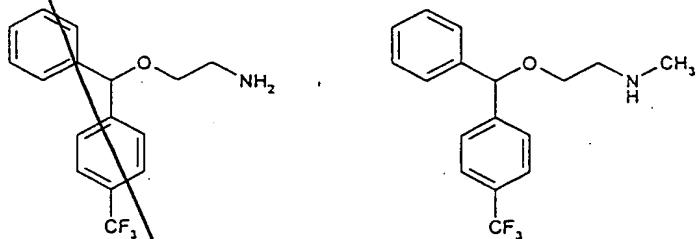
or a pharmaceutically acceptable salt thereof.

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*John C. C.*

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18. A method of treating a patient for depression comprising the step of administering to said patient an effect amount of a compound having the chemical structure:



5 or a pharmaceutically acceptable salt thereof.

*add*  
*137*